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For patients who initially present without advanced or metastatic cancer, an COX-2 inhibitor based drug in combination with another antiangiogenic agent or one or more anticancer agents as an immediate initial therapy
5 prior to surgery, chemotherapy, or radiation therapy, and as a continuous post-treatment therapy in patients at risk for recurrence or metastasis (for example, in adenocarcinoma of the prostate, risk for metastasis is based upon high PSA, high Gleason's score, locally
10 extensive disease, and/or pathological evidence of tumor invasion in the surgical specimen). The goal in these patients is to inhibit the growth of potentially metastatic cells from the primary tumor during surgery or radiotherapy and inhibit the growth of tumor cells
15 from undetectable residual primary tumor.

For patients who initially present with advanced or metastatic cancer, an COX-2 inhibitor based drug in combination with another antiangiogenic agent or one or more anticancer agents of the present invention is used
20 as a continuous supplement to, or possible replacement for hormonal ablation. The goal in these patients is to slow or prevent tumor cell growth from both the untreated primary tumor and from the existing metastatic lesions.

25 In addition, the invention may be particularly efficacious during post-surgical recovery, where the present compositions and methods may be particularly effective in lessening the chances of recurrence of a tumor engendered by shed cells that cannot be removed by
30 surgical intervention.

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Combinations with Other Treatments

The combination of COX-2 inhibitors and antineoplastic agents may be used in conjunction with other treatment modalities, including, but not limited
5 to surgery and radiation, hormonal therapy, antiangiogenic therapy, chemotherapy, immunotherapy, and cryotherapy. The present invention may be used in conjunction with any current or future therapy.

10 The following discussion highlights some agents in this respect, which are illustrative, not limitative. A wide variety of other effective agents also may be used.

Surgery and Radiation

15 In general, surgery and radiation therapy are employed as potentially curative therapies for patients under 70 years of age who present with clinically localized disease and are expected to live at least 10 years.

For example, approximately 70% of newly diagnosed
20 prostate cancer patients fall into this category. Approximately 90% of these patients (65% of total patients) undergo surgery, while approximately 10% of these patients (7% of total patients) undergo radiation therapy. Histopathological examination of surgical specimens reveals that approximately 63% of patients undergoing surgery (40% of total patients) have locally extensive tumors or regional (lymph node) metastasis that was undetected at initial diagnosis. These patients are at a significantly greater risk of recurrence.
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30 Approximately 40% of these patients will actually develop recurrence within five years after surgery.

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Results after radiation are even less encouraging. Approximately 80% of patients who have undergone radiation as their primary therapy have disease persistence or develop recurrence or metastasis within 5 five years after treatment. Currently, most of these surgical and radiotherapy patients generally do not receive any immediate follow-up therapy. Rather, for example, they are monitored frequently for elevated Prostate Specific Antigen ("PSA"), which is the primary 10 indicator of recurrence or metastasis prostate cancer.

Thus, there is considerable opportunity to use the present invention in conjunction with surgical intervention.

15 Hormonal Therapy

Hormonal ablation is the most effective palliative treatment for the 10% of patients presenting with metastatic prostate cancer at initial diagnosis. Hormonal ablation by medication and/or orchietomy is 20 used to block hormones that support the further growth and metastasis of prostate cancer. With time, both the primary and metastatic tumors of virtually all of these patients become hormone-independent and resistant to therapy. Approximately 50% of patients presenting with 25 metastatic disease die within three years after initial diagnosis, and 75% of such patients die within five years after diagnosis. Continuous supplementation with NAALADase inhibitor based drugs are used to prevent or reverse this potentially metastasis-permissive state.

30 Among hormones which may be used in combination with the present inventive compounds, diethylstilbestrol